Scope and Access

The Gene Expression Omnibus (GEO) is a public repository that archives and freely distributes microarray, next-generation sequencing, and other forms of high-throughput functional genomic data submitted by the scientific community. In addition to data storage, a collection of web-based interfaces, browse and search methods, and advanced analytical tools are available to help users find, analyze and download the data stored in GEO. The GEO homepage (https://www.ncbi.nlm.nih.gov/geo/) is the main gateway for searching and browsing high-throughput array data. GEO accepts many categories of high-throughput functional genomic data, including all array-based applications and some high-throughput sequencing data. Questions and issues related to submission should be sent to: geo@ncbi.nlm.nih.gov.

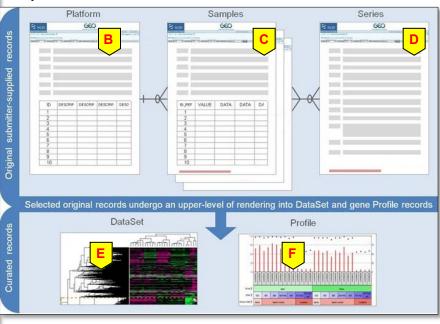
The GEO Databases, Data and Record Types

While GEO was originally established to host gene expression data, it has evolved to host other data types, including comparative genomic analyses, chromatin immunoprecipitation (ChIP) profiling that characterizes genome-protein interactions, non-coding RNA profiling, SNP genotyping and genome methylation status analyses. Thus, the data in GEO is measured in a wide variety of high-throughput assay methods (platforms) and measures a diverse set of sample types, as shown by the GEO Repository Browser (A, https://www.ncbi.nlm.nih.gov/geo/summary/) for the platforms.

Platform, Sample and Series records are submitted to the NCBI by researchers. A Platform record (**B**) is composed of a summary description of the array or sequencer and, for array-based Platforms, a data table defining the array template. A Sample record (**C**) is composed of a description of the biological material, the experimental protocols to which it was subjected, and a data table containing abundance measurements for each feature on the corresponding Platform table. A Series record (**D**) defines a set of related Samples considered to be part of a study, and describes the overall study aim and design.



DataSet records (**E**) are prepared by NCBI Curators with information extracted from the submitter-supplied records. A DataSet represents a summarized collection of consistently processed and experimentally related Sample records categorized according to experimental variables. Profiles (**F**) are derived from DataSets. A Profile consists of the expression measurements for an individual gene across all Samples in a DataSet. DataSets and Profiles are a means for transforming diverse styles of submitted data into a relatively standardized format. These curated database records form the basis of GEO's advanced data display and analysis tools.



Browsing and Searching in GEO

In addition to being able to browse (A) GEO data in sortable, linked tables. GEO Profile and GEO DataSet databases can be searched using query boxes or from their individual homepages linked under the tools column (B). The Entrez search and retrieval system can be used to query for records corresponding to specific GEO accession numbers or for searching with terms for specific data attributes including keywords, authors, organisms, gene symbols, gene names, GenBank accession numbers, or Profiles flagged as being differentially expressed. Users can refine searches by simply typing in terms of interest or constructing very specific fielded que-

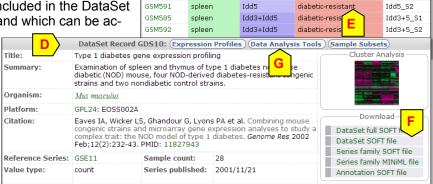


ries. A full listing of all indexed fields, with examples and tips specifically for the GEO Databases, is provided at https://www.ncbi.nlm.nih.gov/geo/info/qqtutorial.html. Additionally, data for large collaborative projects, such as ENCODE and Roadmap Epigenomics, have dedicated browser pages at https://www.ncbi.nlm.nih.gov/geo/info/ENCODE.html and https://www.ncbi.nlm.nih.gov/geo/roadmap/epigenomics/, respectively. Nucleotide or protein sequences can be used to search for genes in GEO Profiles through GEO BLAST (C). This page is also linked from the BLAST homepage (https://blast.ncbi.nlm.nih.gov/) under the "Specialized BLAST" section.

GEO DataSets Summary Pages

The GEO DataSets database stores and displays summary information about each curated experimental set as well as links to the underlying data, including the Platform, Series and other reference material such as organism and PubMed references The DataSet summary page for GDS10 is shown as an example (D). Samples included in the DataSet are linked from a sortable and fully linked table and which can be ac-

cessed with a click of the <u>Sample Subsets</u> button (**E**). Downloading the entire dataset can be accomplished quickly, in many formats (**F**). In addition, due to the curated nature of the data, <u>Data Analytical Tools</u> (**G**) are available for assessment of the data, including statistical comparison and cluster analysis of sample subsets.



tissue

spleen

spleen

spleen

spleen

spleen

GSM582

GSM589

GSM583

GSM590

GSM584

Sample Subsets

Factors

disease state

diabetic-resistant

diabetic-resistant

diabetic-resistant

NOD_S1

NOD S2

Idd3 S1

Idd3 S2

Idd5_S1

diabetic

diabetic

strain

NOD

NOD

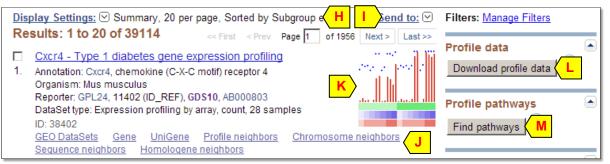
Idd3

Idd3

GEO Profiles Summaries

The GEO Profile database stores gene expression profiles from curated DataSets. Retrieved records are displayed in summary format, which are sortable and downloadable through the <u>Display Setting</u> (H) and <u>Send to</u> (I) links, respectively. The summaries provide information about the particular gene and the Series or DataSet in which it was examined. They are fully linked within GEO and other NCBI databases (J). The Profile Neighbors link and Subgroup effect sort option provide access to other genes that exhibit similar expression behavior. Finally, the chart next to a profile displays the expression level of the gene for comparison of expression levels in all samples within a DataSet (K). Data points for re-

trieved profiles can be downloaded using the "Download profile data" button (L) and pathways involved can be retrieved from Bio-Systems using the "Find pathways" button (M).



ple Subsets

Analyzing with GEO2R

GEO2R is a simple web interface (https://www.ncbi.nlm.nih.gov/geo/geo2r/), which allows sophisticated R-based analysis of GEO data for the identification and visualization of differential gene expression, particularly for those data sets that were not curated by NCBI. Results are presented as a table of genes ordered by significance that can be visualized with GEO Profile graphics. A YouTube video tutorial is available at https://www.youtube.com/watch?v=EUPmGWS8ik0

Customizable GEO DataSet data analysis tools

GEO DataSet records have integrated Data Analysis Tools to facilitate examination and interrogation of the data to help

Compare 2 sets of samples

Find genes

Cluster Analysis

Cluster heatmaps

В

DataSet Record GDS10: Expression Profiles Data Analysis Tools

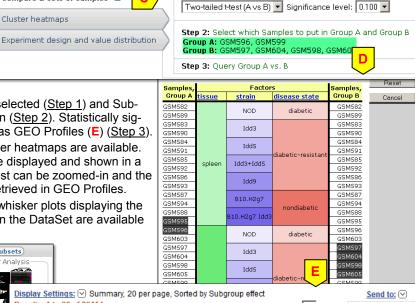
identify potentially interesting genes. The set of tools are accessible from the DataSet Summary page (A) and provides four analysis options:

- Find Genes (B): Querying with gene names or symbols will retrieve expression profiles of those genes examined in the DataSet. Check boxes are made available to display only those flagged as exhibiting differential expression under specific experimental conditions.
- Compare 2 sets of samples (C): Comparisons of data can be performed to retrieve records that are differentially expressed. The type of statistical test
 - to be performed along with significance level can be selected (Step 1) and Subsets of Samples (D) can be chosen for the comparison (Step 2). Statistically significant, differentially expressed genes are displayed as GEO Profiles (E) (Step 3).
- Cluster heatmaps (F): Interactive, customizable, cluster heatmaps are available. Views of several different clustering algorithms can be displayed and shown in a variety of color schemes. In addition, regions of interest can be zoomed-in and the data directly downloaded, displayed in line plots, or retrieved in GEO Profiles.
- Experimental design and value distribution: Box and whisker plots displaying the distribution of expression values of each sample within the DataSet are available to assist with examination of the quality of the data.

DataSet Record GDS10: (Expression Profiles) (Data Analysis Tools) (Sample Subsets)

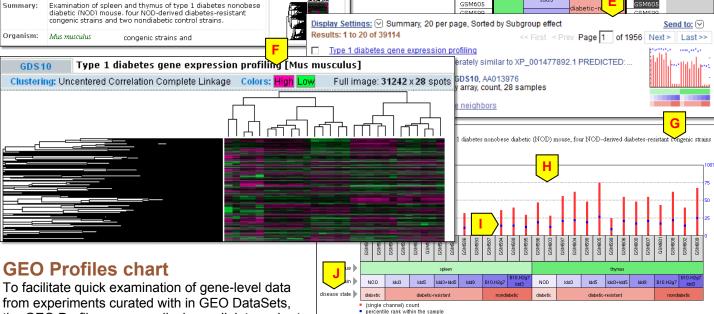
Type 1 diabetes gene expression profiling

Title:



Step 1: Select test and significance level

Data Analysis Tool:



from experiments curated with in GEO DataSets. the GEO Profile summary displays a link to a chart

analysis of gene expression levels. Clicking on the Profiles thumbnail (G) displays a larger version of the chart (H). Full profile details are shown, including descriptions of and links to the DataSet and Samples, as well as the expression profile for the individual gene grouped by experimental subset. The red bars indicate the level of abundance of an individual transcript across the Samples that make up a DataSet, while blue squares indicate rank order for that gene's expression with respect to the other genes within the Sample (I). The colored X-axis bars (J) distinguish named experimental subsets of Samples within the DataSet (type, e.g., 'tissue', description, e.g., 'spleen', and disease state, e.g., 'diabetic'). Located under the chart is a button to display data values, as well as a link to a help document with thorough explanation of this view.

Query DataSets for GSE57926

RBFox2 regulates a broad RNA program and is required for genome-wide

https://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE57926

GEO Submission

GEO accepts many categories of high-throughput functional genomic data, including all array-based applications and some high-throughput sequencing data. The GEO Curation Team is available to assist submitters in depositing their data. Send an E-mail to: geo@ncbi.nlm.nih.gov with a brief description of the type of data you are trying to submit and an explanation of any problems with or questions about the submission procedures. Three different methods for submission available are summarized in the table below.

Submission Method	Format	Key points
GEOarchive	Spreadsheets (e.g., Excel)	Recommended method for most submissions. Quick description of the submitted study or studies using Excel spreadsheet templates.
SOFT	Plain Text	Good option if data and metadata are already in a database which can generate and export data in SOFT plain text format.
MINIML	XML	Good option if data and metadata are already in a database which can generate and export data in MINiML XML format.

All deposit methods described here support the submission of many data types, including: Gene expression, SNP arrays, SAGE, ChIP-chip, Protein arrays, ArrayCGH (Comparative Genomic Hybridization) and high throughput quantitative sequence data. In addition, submissions may remain private until a manuscript describing the data is published. However, as manuscript reviewers often need access to the data for evaluation purposes, GEO has developed a system for creation of a Reviewer URL, providing confidential and anonymous access, which authors can supply in their manuscript submissions. More on GEO submission is available at https://www.ncbi.nlm.nih.gov/geo/info/submission.html.

Downloading Bulk Data Files and Programmatic Access

Data from GEO DataSets and Profiles databases, including all Platform, Sample, Series and DataSet records as well as raw data, are available for bulk download through FTP at ftp://ftp.ncbi.nih.gov/pub/geo/. In addition, metadata from these GEO-specific databases can be accessed programmatically using a suite of programs collectively referred to as the En-

Title

Organism

Relations

BioProject

Series GSE57926

trez Programming Utilities (EUtils). A help page with examples and a link to the complete EUtils documentation is at https://www.ncbi.nlm.nih.gov/geo/info/geo-paccess.html

A Notable Enhancement in Presentation

GEO now makes submitter provided mapping data (in WIG and BEDgraph formats) accessible through the Genome

Data Viewer. Datasets with this enabled are readily retrievable using a fielded query term 'has track"[prop]. Clicking the "See the data on Genome Data Viwer" button (A) of an entry with this enabled brings out this display for that dataset (B).

GEO Documents and References

The collection GEO help documents on data submission, searching, linking, and data analysis are linked from the GEO

SRA SRP042215 Genome Data Viewer See the data on Genome Data Viewer qC1.1 qC1.2 qC2 qC3 qC4 qC5 qD qE1.1 qE2.1 qE2.3 qE3 qE4 qF Pick Assembly ▼ Ideogram View **a** 🔠 https://www.ncbi.nlm.nih.gov/genome/gdv/?context=GEO&acc=GSE57926 3SM1397526 MEFs-SUZ12-NC-ChIPseq NA000084473.1 GSM1397541 Heart-WT-GROseq NA000083900.1 1 1.1 Q- Location, gene or phenotype Your Data

Public on Oct 22, 2015

Mus musculus

targeting of polycomb complex

homepage (https://www.ncbi.nlm.nih.gov/geo) under the "Getting Started" column.

GEO Publications:

- 1. NCBI GEO: archive for functional genomics data sets 10 years on. Barrett T, et.al. Nucleic Acids Res. 2011 Jan;39 (Database issue):D1005-10. Epub 2010 Nov 21. https://www.ncbi.nlm.nih.gov/pubmed/21097893
- 2. NCBI GEO standards and services for microarray data. Edgar R, Barrett T. Nat Biotechnol. 2006 Dec;24(12):1471-2. https://www.ncbi.nlm.nih.gov/pmc/articles/PMC2270403/.
- 3. Gene expression omnibus: microarray data storage, submission, retrieval, and analysis. Barrett T, Edgar R. Methods Enzymol. 2006;411:352-69. https://www.ncbi.nlm.nih.gov/pmc/articles/PMC1619900. See more at https://www.ncbi.nlm.nih.gov/geo/info/citations.html.